

SEQUENCE LISTING

<110> CO, MAN SUNG VASQUEZ, MAXIMILIANO CARRENO, BEATRIZ CELNIKER, ABBIE CHERYL COLLINS, MARY GOLDMAN, SAMUEL GRAY, GARY S. KNIGHT, ANDREA O'HARA, DENISE RUP, BONITA VELDMAN, GEERTRUIDA M. <120> HUMANIZED IMMUNOGLOBULIN REACTIVE WITH B7 MOLECULES AND METHODS OF TREATMENT THEREWITH <130> 08702.0083-00000 <140> 09/501,102 <141> 2000-02-09 <150> 09/339,596 <151> 1999-06-24 <150> 09/249,011 <151> 1999-02-12 <160> 40 <170> PatentIn Ver. 2.1 <210> 1 <211> 405 <212> DNA <213> Mus sp. <220> <221> CDS <222> (1)..(405) <400> 1 atg ggt tgg aac tgt atc atc ttc ttt ctg gtt aca aca gct aca ggt Met Gly Trp Asn Cys Ile Ile Phe Phe Leu Val Thr Thr Ala Thr Gly gtg cac tcc cag gtc cag ctg cag cag tct ggg cct gag ctg gtg agg Val His Ser Gln Val Gln Leu Gln Gln Ser Gly Pro Glu Leu Val Arg cct ggg gaa tca gtg aag att tcc tgc aag ggt tcc ggc tac aca ttc Pro Gly Glu Ser Val Lys Ile Ser Cys Lys Gly Ser Gly Tyr Thr Phe act gat tat gct ata cag tgg gtg aag cag agt cat gca aag agt cta Thr Asp Tyr Ala Ile Gln Trp Val Lys Gln Ser His Ala Lys Ser Leu

	tgg Trp	_		_	_		_			_						240
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	gcc Ala															336
	tac Tyr															384
_	tca Ser 130	_		-												405
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Val	His	Ser	Gln 20	Val	Gln	Leu	Gln	Gln 25	Ser	Gly	Pro	Glu	Leu 30	Val	Arg	
Pro	Gly	Glu 35	Ser	Val	Lys	Ile	Ser 40	Cys	Lys	Gly	Ser	Gly 45	Tyr	Thr	Phe	
Thr	Asp 50	Tyr	Ala	Ile	Gln	Trp 55	Val	Lys	Gln	Ser	His 60	Ala	Lys	Ser	Leu	
Glu 65	Trp	Ile	Gly	Val	Ile 70	Asn	Ile	Туr	Tyr	Asp 75	Asn	Thr	Asn	Tyr	Asn 80	
Gln	Lys	Phe	Lys	Gly 85	Lys	Ala	Thr	Met	Thr 90	Val	Asp	Lys	Ser	Ser 95	Ser	
Thr	Ala	Tyr	Met 100	Glu	Leu	Ala	Arg	Leu 105	Thr	Ser	Glu	Asp	Ser 110	Ala	Ile	
Tyr	Tyr	Cys 115	Ala	Arg	Ala	Ala	Trp 120	Tyr	Met	Asp	Tyr	Trp 125	Gly	Gln	Gly	
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aaa cca ggg cag Lys Pro Gly Gln 65											
gaa tct ggg gtc Glu Ser Gly Val											
ttc act ctc acc Phe Thr Leu Thr 100	atc agc agt Ile Ser Ser	gtg cag gct Val Gln Ala 105	Glu Asp Leu A	ca gtt tat 336 la Val Tyr 10							
tac tgc acg caa Tyr Cys Thr Gln 115	tct tat aat Ser Tyr Asn	ctt tac acg Leu Tyr Thr 120	ttc gga ggg g Phe Gly Gly G 125	gg acc aag 384 ly Thr Lys							
ctg gaa ata aaa Leu Glu Ile Lys 130				396							
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Val Ser Ala Gly Glu Lys Val Thr Met Ser Cys Lys Ser Ser Gln Ser 40 Leu Leu Asn Ser Arg Thr Arg Glu Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr Trp Ala Ser Thr Arg 70 65 Glu Ser Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val Gln Ala Glu Asp Leu Ala Val Tyr 100 Tyr Cys Thr Gln Ser Tyr Asn Leu Tyr Thr Phe Gly Gly Thr Lys 120 125 Leu Glu Ile Lys 130 <210> 5 <211> 405 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: Synthetic nucleotide sequence <220> <221> CDS <222> (1)..(405) atg ggt tgg aac tgt atc atc ttc ttt ctg gtt acc aca gct aca ggt Met Gly Trp Asn Cys Ile Ile Phe Phe Leu Val Thr Thr Ala Thr Gly 5 10 gtg cac tcc cag gtc cag ctg gtg cag tct ggg gct gag gtg aag aag Val His Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys 20 25 cct ggg agc tca gtg aag gtg tcc tgc aaa gct tcc ggc tac aca ttc Pro Gly Ser Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe 35 45 act gat tat gct ata cag tgg gtg aga cag gct cct gga cag ggc ctc Thr Asp Tyr Ala Ile Gln Trp Val Arg Gln Ala Pro Gly Gln Gly Leu 50 55 gag tgg att gga gtt att aat att tac tat gat aat aca aac tac aac Glu Trp Ile Gly Val Ile Asn Ile Tyr Tyr Asp Asn Thr Asn Tyr Asn

cag aag ttt aag ggc aag gcc aca atg act gta gac aag tcg acg agc 288 Gln Lys Phe Lys Gly Lys Ala Thr Met Thr Val Asp Lys Ser Thr Ser 85 90 aca gcc tat atg gaa ctt agt tct ttg aga tct gag gat acg gcc gtt 336 Thr Ala Tyr Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val 100 105 110 tat tac tgt gca aga gcg gcc tgg tat atg gac tac tgg ggt caa ggt 384 Tyr Tyr Cys Ala Arg Ala Ala Trp Tyr Met Asp Tyr Trp Gly Gln Gly 120 405 acc ctt gtc acc gtc tcc tca Thr Leu Val Thr Val Ser Ser 130 <210> 6 <211> 135 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence: Synthetic peptide sequence <400> 6 Met Gly Trp Asn Cys Ile Ile Phe Phe Leu Val Thr Thr Ala Thr Gly 10 Val His Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ser Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe 40 Thr Asp Tyr Ala Ile Gln Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Ile Gly Val Ile Asn Ile Tyr Tyr Asp Asn Thr Asn Tyr Asn Gln Lys Phe Lys Gly Lys Ala Thr Met Thr Val Asp Lys Ser Thr Ser 85 Thr Ala Tyr Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val 105 Tyr Tyr Cys Ala Arg Ala Ala Trp Tyr Met Asp Tyr Trp Gly Gln Gly 120 Thr Leu Val Thr Val Ser Ser 130

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                                     10
ggc acc tgt ggg gac att gtg ctg aca cag tct cca gat tcc ctg gct
Gly Thr Cys Gly Asp Ile Val Leu Thr Gln Ser Pro Asp Ser Leu Ala
gta agc tta gga gag agg gcc act att agc tgc aaa tcc agt cag agt
Val Ser Leu Gly Glu Arg Ala Thr Ile Ser Cys Lys Ser Ser Gln Ser
         35
ctg ctc aac agt aga acc cga gag aac tac ttg gct tgg tac cag cag
Leu Leu Asn Ser Arg Thr Arg Glu Asn Tyr Leu Ala Trp Tyr Gln Gln
     50
                         55
aaa cca ggg cag cct cct aaa ctg ctg atc tac tgg gca tcc act agg
Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Trp Ala Ser Thr Arg
 65
                     70
gaa tot ggg gtc cot gat cgc ttc agt ggc agt gga tot ggg aca gat
                                                                   288
Glu Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp
                 85
                                     90
ttc act ctc acc atc agc agt ctg cag gct gaa gac gtg gca gtt tat
                                                                   336
Phe Thr Leu Thr Ile Ser Ser Leu Gln Ala Glu Asp Val Ala Val Tyr
            100
                                105
tac tgc acg caa tct tat aat ctt tac acg ttc gga cag ggg acc aag
                                                                   384
Tyr Cys Thr Gln Ser Tyr Asn Leu Tyr Thr Phe Gly Gln Gly Thr Lys
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                            120
gtg gaa ata aaa
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Val Glu Ile Lys
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Gly Thr Cys Gly Asp Ile Val Leu Thr Gln Ser Pro Asp Ser Leu Ala
Val Ser Leu Gly Glu Arg Ala Thr Ile Ser Cys Lys Ser Ser Gln Ser
                             40
Leu Leu Asn Ser Arg Thr Arg Glu Asn Tyr Leu Ala Trp Tyr Gln Gln
Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Trp Ala Ser Thr Arg
Glu Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp
                                     90
Phe Thr Leu Thr Ile Ser Ser Leu Gln Ala Glu Asp Val Ala Val Tyr
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Val Glu Ile Lys
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Val Ile Asn Ile Tyr Tyr Asp Asn Thr Asn Tyr Asn Gln Lys Phe Lys
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ggc
Gly
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                                                                   51
gct
Ala
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gtc aat tca gag gtt cac ctg cag cag tct ggg gct gag ctt gtg agg
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Val Asn Ser Glu Val His Leu Gln Gln Ser Gly Ala Glu Leu Val Arg
            2.0
                                 25
cca ggg gcc tta gtc aag ttg tcc tgc aaa cct tct ggc ttc aac att
Pro Gly Ala Leu Val Lys Leu Ser Cys Lys Pro Ser Gly Phe Asn Ile
                            40
aaa gac tac tat atg cac tgg gtg aag cag agg cct gaa cag ggc ctg
Lys Asp Tyr Tyr Met His Trp Val Lys Gln Arg Pro Glu Gln Gly Leu
   50
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55

					att Ile 70	-										240
					aag Lys											288
	_				ctc Leu											336
				-	gag Glu					_						384
_	_	_	_	_	tct Ser											405
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Val	Asn	Ser	Glu 20	Val	His	Leu	Gln	Gln 25	Ser	Gly	Ala	Glu	Leu 30	Val	Arg	
Pro	Gly	Ala 35	Leu	Val	Lys	Leu		Cys	Lvs	Pro	Ser	Clv	Phe	λen	Ile	
Lys	_						40		-2-			45	1110	ABII		
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Glu 65	50				His Ile 70	55	Val		Gln	Arg	Pro 60	45 Glu	Gln	Gly		
65	50 Trp	Ile	Gly	Trp	Ile	55 Asp	Val Pro	Glu	Gln Asn	Arg Gly 75	Pro 60 Asn	45 Glu Thr	Gln Leu	Gly Tyr	qaA 08	
65 Pro	50 Trp Lys	Ile Phe	Gly Gln	Trp Gly 85	Ile 70	55 Asp Ala	Val Pro Ser	Glu Ile	Gln Asn Thr 90	Arg Gly 75 Ala	Pro 60 Asn Asp	45 Glu Thr	Gln Leu Ser	Gly Tyr Ser 95	Asp 80 Asn	
65 Pro Thr	50 Trp Lys Ala	Ile Phe Tyr	Gly Gln Leu 100	Trp Gly 85 Gln	Ile 70 Lys	55 Asp Ala Ser	Val Pro Ser Ser	Glu Ile Leu 105	Gln Asn Thr 90 Thr	Arg Gly 75 Ala	Pro 60 Asn Asp Glu	45 Glu Thr Thr	Gln Leu Ser Thr	Gly Tyr Ser 95 Ala	Asp 80 Asn Val	

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gtc ata ttg tcc agt gga gaa att gtg ctc acc cag tct cca gca ctc
Val Ile Leu Ser Ser Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Leu
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atg gct gca tct cca ggg gag aag gtc acc atc acc tgc agt gtc agc
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Met Ala Ala Ser Pro Gly Glu Lys Val Thr Ile Thr Cys Ser Val Ser
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                             4.0
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Ser Ser Ile Ser Ser Ser Asn Leu His Trp Tyr Gln Gln Lys Ser Glu
     50
acc tcc ccc aaa ccc tgg att tat ggc aca tcc aac ctg gct tct gga
                                                                   240
Thr Ser Pro Lys Pro Trp Ile Tyr Gly Thr Ser Asn Leu Ala Ser Gly
65
                     70
gtc cct gtt cgc ttc agt ggc agt gga tct ggg acc tct tat tct ctc
Val Pro Val Arg Phe Ser Gly Ser Gly Ser Gly Thr Ser Tyr Ser Leu
aca atc agc agc atg gag gct gaa gat gct gcc act tat tac tgt caa
Thr Ile Ser Ser Met Glu Ala Glu Asp Ala Ala Thr Tyr Tyr Cys Gln
            100
cag tgg agt agt tac cca ctc acg ttc ggt gct ggg acc aag ctg gag
Gln Trp Ser Ser Tyr Pro Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu
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Leu Lys
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Val Ile Leu Ser Ser Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Leu
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25

Met Ala Ala Ser Pro Gly Glu Lys Val Thr Ile Thr Cys Ser Val Ser 4.0 3.5 Ser Ser Ile Ser Ser Ser Asn Leu His Trp Tyr Gln Gln Lys Ser Glu Thr Ser Pro Lys Pro Trp Ile Tyr Gly Thr Ser Asn Leu Ala Ser Gly 70 75 Val Pro Val Arg Phe Ser Gly Ser Gly Ser Gly Thr Ser Tyr Ser Leu Thr Ile Ser Ser Met Glu Ala Glu Asp Ala Ala Thr Tyr Tyr Cys Gln 105 100 Gln Trp Ser Ser Tyr Pro Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu 120 125 Leu Lys 130 <210> 25 <211> 405 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: Synthetic nucleotide sequence <220> <221> CDS <222> (1)..(405) atg aaa tgc agc tgg gtc atc ttc ttc ctg atg gca gtg gtt aca ggg Met Lys Cys Ser Trp Val Ile Phe Phe Leu Met Ala Val Val Thr Gly 10 gtc aat tca gag gtt cag ctg gtg cag tct ggg gct gag gtt aag aag Val Asn Ser Glu Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys 20 25 cca ggg gcc tca gtc aag gtg tcc tgc aaa cct tct ggc ttc aac att Pro Gly Ala Ser Val Lys Val Ser Cys Lys Pro Ser Gly Phe Asn Ile 35 40 45 aaa gac tac tat atg cac tgg gtg agg cag gcg cct gga cag ggc ctc Lys Asp Tyr Tyr Met His Trp Val Arg Gln Ala Pro Gly Gln Gly Leu 50 55 gag tgg att gga tgg att gat cct gag aat ggt aat act cta tat gac Glu Trp Ile Gly Trp Ile Asp Pro Glu Asn Gly Asn Thr Leu Tyr Asp 75 70

ccg aag Pro Lys													288
aca gcc Thr Ala											_		336
tat tac Tyr Tyr													384
acc ctg Thr Leu 130	_												405
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Pro Gly	Ala Ser 35	Val Ly	s Val	Ser 40	Суѕ	Lys	Pro	Ser	Gly 45	Phe	Asn	Ile	
Lys Asp 50	Tyr Tyr	Met H	s Trp 55	Val	Arg	Gln	Ala	Pro 60	Gly	Gln	Gly	Leu	
Glu Trp 65	Ile Gly		e Asp	Pro	Glu	Asn	Gly 75	Asn	Thr	Leu	Tyr	Asp 80	
Pro Lys	Phe Gln	Gly Ly 85	s Ala	Thr	Ile	Thr 90	Ala	Asp	Thr	Ser	Thr 95	Ser	
Thr Ala	Tyr Met 100	Glu Le	eu Ser	Ser	Leu 105	Arg	Ser	Glu	Asp	Thr 110	Ala	Val	
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Thr Leu 130	Val Thr	Val Se	er Ser 135										

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Val Ile Leu Ser Ser Gly Asp Ile Gln Met Thr Gln Ser Pro Ser Ser
                                 25
ctg tct gca tct gta ggg gat agg gtc acc atc acc tgc agt gtc agc
Leu Ser Ala Ser Val Gly Asp Arg Val Thr Ile Thr Cys Ser Val Ser
         35
                             40
tca agt ata agt tcc agc aac ttg cac tgg tac cag cag aag cca ggc
                                                                   192
Ser Ser Ile Ser Ser Ser Asn Leu His Trp Tyr Gln Gln Lys Pro Gly
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                         55
aag goo coo aaa coo ttg att tat ggo aca too aac ctg got tot gga
                                                                   240
Lys Ala Pro Lys Pro Leu Ile Tyr Gly Thr Ser Asn Leu Ala Ser Gly
 65
                     70
gtc cct agt cgc ttc agt ggc agt gga tct ggg acc gat tat act ctc
                                                                   288
Val Pro Ser Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Tyr Thr Leu
                 85
aca atc agc agc ttg cag cct gaa gat gtt gcc act tat tac tgt caa
                                                                   336
Thr Ile Ser Ser Leu Gln Pro Glu Asp Val Ala Thr Tyr Tyr Cys Gln
cag tgg agt agt tac cca ctc acg ttc ggt caa ggg acc aag gtg gag
                                                                   384
Gln Trp Ser Ser Tyr Pro Leu Thr Phe Gly Gln Gly Thr Lys Val Glu
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                            120
atc aaa
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Ile Lys
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peptide sequence

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Leu Ser Ala Ser Val Gly Asp Arg Val Thr Ile Thr Cys Ser Val Ser
Ser Ser Ile Ser Ser Ser Asn Leu His Trp Tyr Gln Gln Lys Pro Gly
                         55
    50
Lys Ala Pro Lys Pro Leu Ile Tyr Gly Thr Ser Asn Leu Ala Ser Gly
Val Pro Ser Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Tyr Thr Leu
Thr Ile Ser Ser Leu Gln Pro Glu Asp Val Ala Thr Tyr Tyr Cys Gln
Gln Trp Ser Ser Tyr Pro Leu Thr Phe Gly Gln Gly Thr Lys Val Glu
Ile Lys
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                                                                 21
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Gly Thr Ser Asn Leu Ala Ser
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Gln Gln Trp Ser Ser Tyr Pro Leu Thr
1
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Gln Gln Trp Ser Ser Tyr Pro Leu Thr
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Asp Tyr Tyr Met His
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                  5
                                     10
ggc
                                                                   51
Gly
<210> 38
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<400> 38
Trp Ile Asp Pro Glu Asn Gly Asn Thr Leu Tyr Asp Pro Lys Phe Gln
                5
                                    10
Gly
<210> 39
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